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include Cisplatin, actinomycin D, 5-fluorouracil, bleomycin, and cyclophosphamide methotrxate.

In the treatment of superficial bladder cancer, COX-2 inhibitors can be used to treat the disease in combination with other COX-2 inhibitors, or in combination with surgery (TUR), chemotherapy and intravesical therapies.

A preferred therapy for the treatment of superficial bladder cancer is a combination of therapeutically effective amounts of one or more COX-2 inhibitors in combination with: thiotepa (30 to 60 mg/day), mitomycin C (20 to 60 mg/day), and doxorubicin (20 to 80 mg/day).

A preferred intravesicle immunotherapeutic agent
that may be used in the present invention is BCG. A
preferred daily dose ranges from 60 to 120 mg, depending
on the strain of the live attenuated tuberculosis
organism used.

A preferred photodynamic therapuetic agent that may

be used with the present invention is Photofrin I, a

photosensitizing agent, administered intravenously. It

is taken up by the low-density lipoprotein receptors of

the tumor cells and is activated by exposure to visible

light. Additionally, neomydium YAG laser activation

generates large amounts of cytotoxic free radicals and

singlet oxygen.

In the treatment of muscle-invasive bladder cancer, COX-2 inhibitors can be used to treat the disease in combination with other COX-2 inhibitors, or in combination with surgery (TUR), intravesical

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chemotherapy, radiation therapy, and radical cystectomy with pelvic lymph node dissection.

A preferred radiation dose for the treatment of bladder cancer is between 5,000 to 7,000 cGY in 5 fractions of 180 to 200 cGY to the tumor. Additionally, 3,500 to 4,700 cGY total dose is administered to the normal bladder and pelvic contents in a four-field technique. Radiation therapy should be considered only if the patient is not a surgical candidate, but may be considered as preoperative therapy.

A preferred combination of surgery and chemotherapeutic agents that can be used in combination with the COX-2 inhibitors of the present invention is cystectomy in conjunction with five cycles of cisplatin (70 to 100 mg/m(square)); doxorubicin (50 to 60 mg/m(square); and cyclophosphamide (500 to 600 mg/m(square).

A more preferred therapy for the treatment of superficial bladder cancer is a combination of therapeutically effective amounts of one or more COX-2 inhibitors.

An even more preferred combination for the treatment of superficial bladder cancer is a combination of therapeutically effective amounts of one or more COX-2 inhibitors in combination with the following combinations of antineoplastic agents: 1) cisplatin, doxorubicin, cyclophosphamide; and 2) cisplatin, 5fluorouracil. An even more preferred combination of chemotherapeutic agents that can be used in combination with radiation therapy and the COX-2 inhibitors is a combination of cisplatin, methotrexate, vinblastine.

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Currently no curative therapy exists for metastatic bladder cancer. The present invention contemplates an effective treatment of bladder cancer leading to improved tumor inhibition or regression, as compared to current therapies.

In the treatment of metastatic bladder cancer, COX-2 inhibitors can be used to treat the disease in combination with other antiangiogenic agents, or in combination with surgery, radiation therapy or with chemotherapeutic agents.

A preferred therapy for the treatment of metastatic bladder cancer is a combination of therapeutically effective amounts of one or more COX-2 inhibitors.

A more preferred combination for the treatment of

metastatic bladder caner is a combination of
therapeutically effective amounts of one or more COX-2
inhibitors in combination with the following
combinations of antineoplasitc agents: 1) cisplatin and
methotrexate; 2) doxorubicin, vinblastine,

20 cyclophoshamide, and 5-fluorouracil; 3) vinblastine, doxorubicin, cisplatin, methotrexate; 4) vinblastine, cisplatin, methotrexate; 5) cyclophosphamide, doxorubicin, cisplatin; 6) 5-fluorouracil, cisplatin.

25 <u>Example 6</u>

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Pancreas Cancer

Approximately 2% of new cancer cases diagnoses in the United States is pancreatic cancer. Pancreatic cancer is generally classified into two clinical types: 1) adenocarcinoma (metastatic and non-metastatic), and